Amendment to the Drawings:

The attached sheets of drawings – both "Replacement" and "Annotated" sheets – represent Figures 3, 4, 7, 17, 19, 20, and 22 amended as required in the Office Action.

Figures 3, 19, and 22 are amended to separately identify Fig. 3A, Fig. 3B, Fig. 19A, Fig. 19B, Fig. 22A, and Fig. 22B.

Figures 4, 7, 17, and 20 are amended to identify every sequence in the figures with the corresponding SEQ ID NO:, as found in the "Sequence Listing."

Replacement Sheets of Proposed Drawings Corrections

Application Number 09/787,443 Amendment September 15, 2003 Replacement Sheet 3/25

PCT/DK99/00500

SEP 1 5 2003 27 15

Stimulation of neurite outgrowth by the C3-peptide.

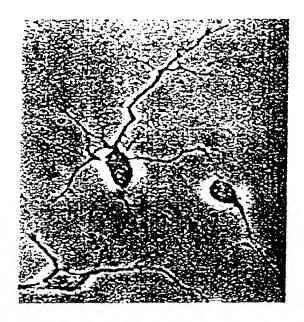


Fig. 3B

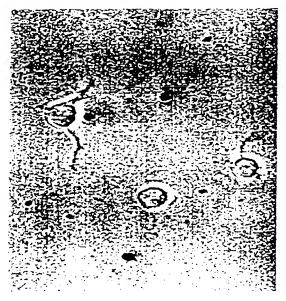


Fig. 3A

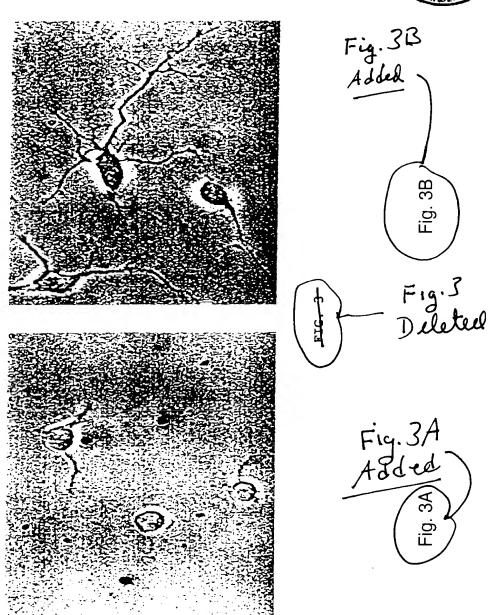
Application Number 09/787,443 Amendment September 15, 2003

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PCT/DK99/00500

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Stimulation of neurite outgrowth by the C3-peptide.



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NCAM-Ig1 binding sequences identified from a combinatorial library of synthetic peptides.

| A | A R A L N W G A K P K A G S A V K L K K K A A K Y V L L P L R L S A S T K R S M Q G L - A R R A L L V R V N R L A Y Y L L V R V N R L A Y Y L L V R V N R L A K R S W G P L L N R L A K R S V Q K L D G Q A R Q K T M K P R R S A G D Y N P D L D R - A S K K P K R N L D G Q A R Q K T M K P R R S A G D Y N P D L D R - A S K K P K R N L K K A A K K E K P K R N L T K K A K K E K P N K P N D A Q M G R Q S L D R N A E G G K K K K M R A A K K E R Q R K D T Q A K K K E R Q R K D T Q A K K K E R Q R K D T Q A K K S R K G N S S L M A R K S R D M T A L K | SEQ ID NO.: 3 4 5 6 7 8 9 10 11 12 13 1 14 15 16 17 18 19 2 20 21 22 |
|-------------|---|---|
| B C3 | A S K K P K R N I K A A K R N G P L I N R I A K R S V Q K L D G Q A S T K R S M Q G I - K T G R R P R N W G A K P K Q K T M K P R R S A K E R Q R K D T Q A K K E K P N K P N D | 39 40 41 42 43 44 |
| D D4 | ARKTKSRERKD ARKTKSRERKD ARKTKSRERKD | 3 47 |

FIG. 4

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Annotated Marked-up Sheet

PCT/DK99/00500

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NCAM-Ig1 binding sequences identified from a combinatorial library of synthetic peptides.



| | | | | | | | | | | | | | | | | | , | A | |
|----|-----------------------------|--|------------------------|--------------------|------------------------|------------------------|----------------------------|----------------------------|------------------------|---------------------|-------------------------|------------------------|--------------|-------|--------|----|----|--|--|
| A | | .\ .\ .\ .\ .\ .\ .\ .\ .\ | RGKSRYTKKRGSRSPKQEKKKR | SYTRYNRRODKKOKKMG | LAVKALKNSKYKTALEGGEKRS | SVERTIKGVTNURKDKRKREKR | KISLVTPQMPKERRPQKQQG | 22222 | AKTQUERLEBLNKKLKLKKQST | K R G | PRILARPRORRKKPKNRRTNLL | KAS-LIRIOS-ADRKDBAQAMK | | | | | SI | EQ ID NO.: 3 4 5 6 7 8 9 10 11 12 13 1 14 15 16 17 18 19 2 20 21 22 | |
| | C3 T N K R A L A R | K | S T W K | G G | K A R A M | S R K | K A A T P P | R K K K R K | N R R R | | <u>K</u> G V M | <u>A</u> P Q Q | I. K G | Z 🗅 . | R G | Ĭ. | | 39 40 41 42 43 44 | |
| C. | 153 | <u>۸</u> ۸ | | <u>К</u> К К | E E T | R K K | Q P S | R N R | K K E | <u>D</u> I' R | T N K | n n | • | | | | | 2 45 46 | |
| D | D4 | <u>\</u> | <u>R</u> T | Z Z | <u>L</u> K | 7 7 | W T | <u>C</u> | K K | K R | P P | K R | | | | | | 3 47 | |

FIG. 4

Column idenlifying a SFO ID NO: for each sequence Added.

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SEP 1 5 2003 22

Effect of NCAM-Ig1 binding peptides on cell aggregation and neurite outgrowth. $\,$

WO 00/18801

Controls for NCAM Ig1 binding peptide (C3)

| | l'eptide. | | | | | Sequence | 444 | | | | | | Effect | . 1 |
|---------------|----------------------|------------|---------------|--|----------|----------|------------|------------|------------|------------|------------|--------------|---------------|---------------|
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| 2 | C.Macetyl. K (120) / | 70) A | S | ¥ | # * | _ | ž | 4 | <u>.</u> | | į | : | | |
| | Ala subst K/R | | | | | | : | • | 7 | _ | > | 4 | ‡ | ! |
| c | 116 | < | S | × | × | ۵. | ¥ | < | z : | | ٤ : | ٠ ، | | • |
| ე <u>-</u> | 211 | . ∢ | S | ¥ | ¥ | ۵. | ~ | ~ | z | _ | 4 : | < ∙ | > ' | > • |
| 4 1 | | < < | | > | < | ۵. | ~ | < | z | | × | < | - | 9 |
| 2 | 8 | < ⋅ | י נ | ٠ - | : - | . α | ~ | ~ | z | _ | × | < | 0 | 0 |
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.IG. 7

acetylation on lysine

| 11. – SEQ ID NO.: 2 | 12. – SEQ ID NO.: 36 | 13. – SEQ ID NO.: 3 | 14. – SEQ ID NO.: 37 | 15. – SEQ ID NO.: 38 |
|---------------------|----------------------|---------------------|----------------------|----------------------|
| 6. – SEQ ID NO.: 31 | 7. – SEQ ID NO.: 32 | 8. – SEQ ID NO.: 33 | 9. – SEQ ID NO.: 34 | 10. – SEQ ID NO.: 35 |
| 1 SEQ ID NO.: 1 | 2. – SEQ ID NO.: 27 | 3 SFQ ID NO.: 28 | 4. – SEQ ID NO.: 29 | 5 - SFO ID NO.: 30 |

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Annotated Marked-up Sheet

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PCT/DK99/00500

Effect of NCAM-Ig1 binding peptides on cell aggregation and neurite outgrowth.

| l'epijde | | | | | Sequence | 44 | | | | | | Effect | |
|---|------------|-----------|-----------|------------|------------|------------|------------|------------|---------------|------------|------------|-------------|----------|
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| Ala subst K/R | | | | | i | ; | | ; | _ | > | • | 1 | ! |
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| Scranded C3 | | | , | ; | 2 | 2 | _ | v | 4 | z | ۵ | ‡ | : |
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| 9 114 | c. | Z | ≺ | s i | _ = | × • | 4 د | ۷ ــ | د د | : ◄ | : × | ‡ | : |
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| 14 Scrumbled D4 | ပ | د | × | æ | ≱ | < | ٠, | Σ. | ۷ | c | : | | |
| Poly-K | : | 2 | ۷ | 7 | × | × | | | | | | • | |
| 15 K6 (dendrimer.11 | × | 4 | ۷ | ź | : | ; | | | | | | | |
| • effect on neurite extension (neur) and aggregation (agg | rite exter | ısion (ne | ur) and a | ggregatic | in (agg) | | | | | | | | |

the corresponding SEQIDNO: Added

11. - SEQ ID NO.: 2

12. – SEQ ID NO.: 36 13. – SEQ ID NO.: 3 14. – SEQ ID NO.: 37 15. – SEQ ID NO.: 38 - SEQ ID NO.: 31 - SEQ ID NO.: 32 - SEQ ID NO.: 33 - SEQ ID NO.: 34 - SEQ ID NO.: 35

- SEQ ID NO.: 28 - SEQ ID NO.: 29 - SEQ ID NO.: 30

- SEQ ID NO.: 27

acetylation on lysine

SUSBSTITUTE SHEET (RULE 26)

Controls for NCAM Ig1 binding peptide (C3)

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17/25

The predicted amino acid sequence of human NCAM-140. (SEQ ID NO: 48)



VRITEERIPN HOGGKHIEPN EITPLIEPEK CPVEAKPECO EIETKPAPAE VRIVPNDATO EPSAPKLEGQ MGEDGNSIKV NLIKQDDGGS PIRHYLVRYR ALSSEWKPEI RLPSGSDHVM GEEVWHSKWY DAKEASMEGI VTIVGIKPET TYAVRIAALN GKGIGEISAA SEFKTOPVQG LKSLDWNAEY EVYVVAENOO GKSKAAHFVF RTSBOPTAIP ANGSPTSGLS TGALVGILLV IFVLLLUVVD ITCYFLNKCG LFMCIAVNLC GKAGPGAKGK DMERGKAAPS KDRSKRPIVE VRSHARVSSL TLKSIQYTDA GEYICTASNT IGQDSQSMYL EVQYAPKLQG PVAVYTWEGN AVNRIGQESL EFILVQADTP SSPSIDQVEP YSSTAQVQFD EPEATGGVPI LKYKAEWRAV KNAPTPQEFR EGEDAVIVCD VVSSLPPTII WKHKGRDVIL KXDVRFIVLS NNYLQIRGIK KTDEGTYRCE GRILARGEIN PKDIQVIVNV PPTIQARQNI VNATANLGQS VTLVCDAEGF 301 HIKVFAKPKI TYVENQTAME LEBQVTLTCE ASGDPIPSIT WRTSTRNISS EEKTLDGHAV QUNITCEVFA YPSATISWFR DGQLLPSSNY SNIKIYNTPS ASYLEVTPDS ENDFGNYNCT PEPTMSWIKD GEQIEQEEDD EKYIPSDDSS QLIIKKVDKN DEAEYICIAE NKAGEQDAII 61 KLTPNQQRIS VVWNDDSSST LTIYNANIDD AGIYKCVVTG EDGSESEATV NVKIPQKLMF 1 MIQTKOLIWT LFFLGTAVSL QVDIVPSQGE ISVGESKFFL CQVAGDAKDK DISWFSPNGE 481 541 121 241 361

16.17

841 TKENESKA

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KNAPTPQEFR EGEDAVIVCD VVSSLPPTII WKHKGRDVIL KXDVRFIVLS NNYLQIRGIK

121

61 KLTPNQQRIS VVWNDDSSST LTIYNANIDD AGIYKCVVTG EDGSESEATV NVKIPQKLMF

1 MLQTKOLIWT LFFLGTAVSL QVDIVPSQGE ISVGESKFFL CQVAGDAKDK DISWFSPNGE

Annotated Marked-up Sheet

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PCT/DK99/00500 O I P

The predicted amino acid sequence of human NCAM-140.

(SEQ ID NO: 48)

SEQ IDNO: identifying the sequence Added

721 IFVLLLVVVD ITCYFLNKCG LFMCIAVNIC GKAGPGAKGK DMERGKAAPS KDRSKEPIVE VRIEEERIPN HOGGKHIEPN EITPLIEPEK CPVEAKPECO EIETKPAPAE VKIVPNDAIO EPSAPKLEGO MGEDGNSIKV NLIKODDGGS PIRHYLVRYR ALSSEWKPEI RLPSGSDHVM 481 AVNRIGQESL EFILVQADTP SSPSIDQVEP YSSTAQVQFD EPEATGGVPI LKYKAEWRAV 301 HLKVPAKPKI TYVENQTAME LEBQVTLTCE ASGDPIPSIT WRTSTRNISS EEKTLDGHMV VRSHARVSSL TLKSIQYTDA GEYICTASNT IGQDSQSMYL EVQYAPKLQG PVAVYTWEGN 421 QVNITCEVFA YPSATISWPR DGQLLPSSNY SNIKIYNTPS ASYLEVTPDS ENDFGNYNCT 541 GEEVWHSKWY DAKEASWEGI VTIVGLKPET TYAVRLAALN GKGLGEISAA SEFKTOPVOG LKSLDWNARY EVYVVAENOO GKSKAAHFVF RTSBOPTAIP ANGSPTSGLS TGAIVGILIV PEPTMSWIKD GEQIEQEEDD EKYIPSDDSS QLTIKKVDKN DEAEYICIAE NKAGEQDATI KIDEGIYRCE GRILARGEIN PKDIQVIVNV PPTIQARQNI VNATANIGQS VTLVCDAEGF TKENESKA 181 361 241

F16.1

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The effect of the NCAM Ig2 domain and the Ig2-p peptide and control peptides derived from the Ig2-p peptide on cell aggregation.

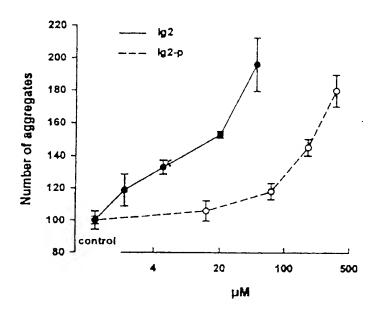


Fig. 19A

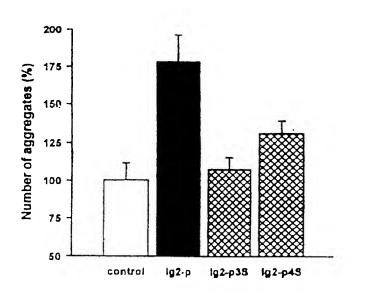


Fig. 19B

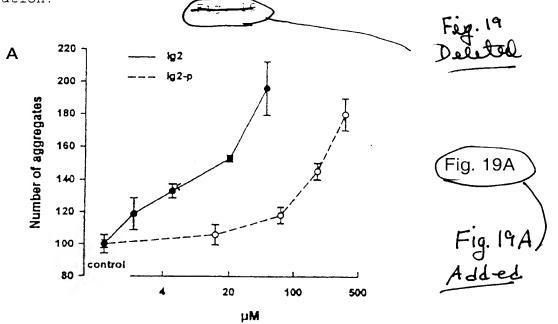
Application Number 09/787,443 WO 00/18801

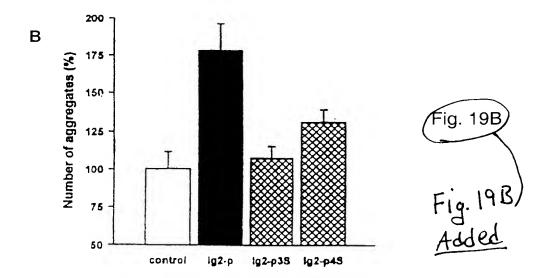
Amendment September 15, 2003

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The effect of the NCAM Ig2 domain and the Ig2-p peptide and control peptides derived from the Ig2-p peptide on cell aggregation.





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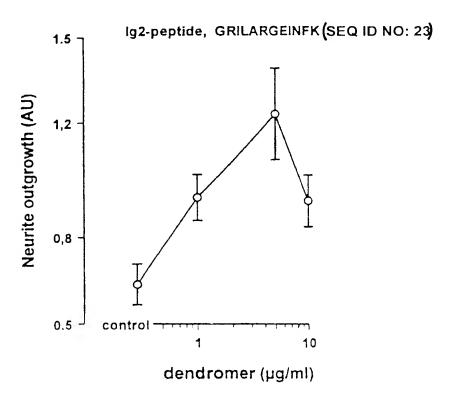
Replacement Sheet

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The effect of the Ig2-p peptide dendrimer on neurite outgrowth.

Fig. 20



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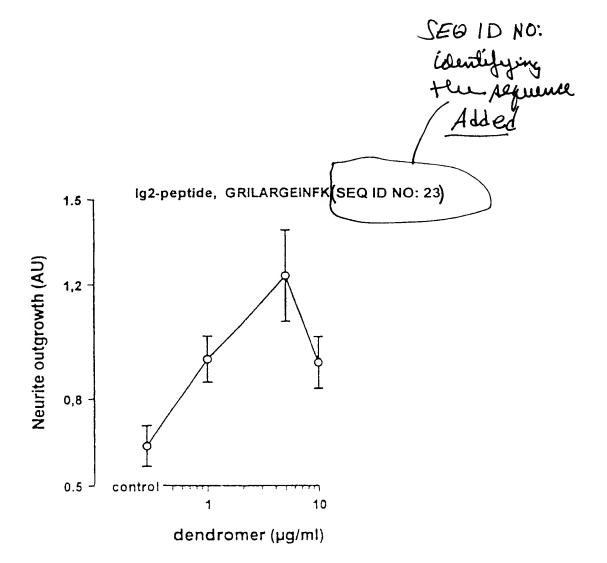
PCT/DK99/00500

20/25

The effect of the Ig2-p peptide dendrimer on neurite outgrowth.

Fig. 20





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Micrograph showing the effect of the Ig2-p peptide on neurite outgrowth.

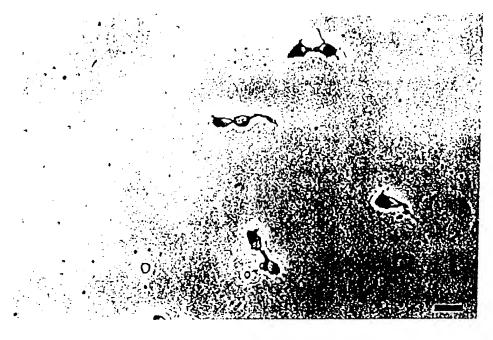


Fig. 22A

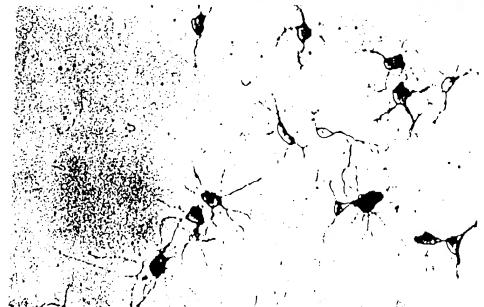


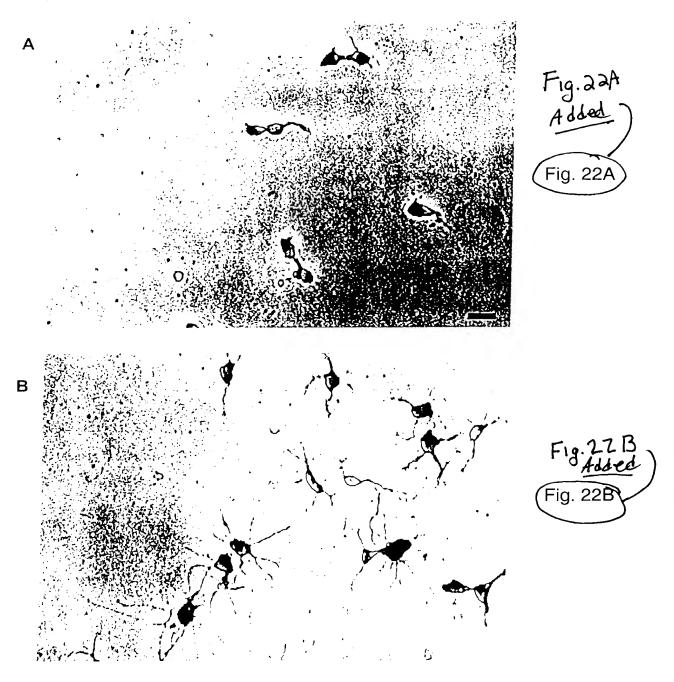
Fig. 22B

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Micrograph showing the effect of the $\ensuremath{\operatorname{Ig2-p}}$ peptide on neurite outgrowth.



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